

FIG. 1A

```

1  GTCTTCCACCATGCACTCGCTGGGCTTCTCTCTGTGGCGTGTCTCTGCTGCCCGCTG 60
   -----+-----+-----+-----+-----+-----+-----+-----+
   CAGGAAGGTGGTACGTAGCGACCCGAAGAAGAGACACCGCACAAAGAGACGAGCGGCGAC
       M H S L G F F S V A C S L L A A A -
C
61  CGCTGCTCCCGGGTCTCGCGAGGGCGCCCGCGCGCGCGCTTCGAGTCCGGACTCG 120
   -----+-----+-----+-----+-----+-----+-----+-----+
   GCGACGAGGGCCAGGAGCGCTCCGCGGGCGGGCGGGCGGAGCTCAGGCCCTGAGC
       L L P G P R E A P A A A A A A F E S G L D -
C
121 ACCTCTCGGACGGGAGCCCGACCGGGCGAGGCCACGGCTTATGCAAGCAAAGATCTGG 180
   -----+-----+-----+-----+-----+-----+-----+-----+
   TGGAGAGCCTGCGCCTCGGGCTGCGCCCGCTCCGGTGCCGAATACGTTCTTCTAGACC
       L S D A E P D A G E A T A Y A S K D L E -
C
181 AGGAGCAGTTACGGTCTGTGTCCAGTGTAGATGAACCTCATGACTGTACTTACCCAGAAT 240
   -----+-----+-----+-----+-----+-----+-----+-----+
   TCCTCGTCAATGCCAGACACAGGTCACATCTACTTGAGTACTGACATGAGATGGGTCCTTA
       E Q L R S V S S V D E L M T V L Y P E Y -
C
241 ATTGGAAAATGTACAAGTGTACAGTCTAGGAAAGGAGGCTGGCAACATAACAGAGAACAGG 300
   -----+-----+-----+-----+-----+-----+-----+-----+
   TAACCTTTTACATGTTACACAGTCGATTCCTTTCTCCGACCGTTGTATGTTCTTGTCC
       W K M Y K C Q L R K G G W Q H N R E Q A -
C
   CCAACCTCAACTCAAGGACAGAGACTATAAAATTTGCTGCAGCACATTATAATACAG

```

MATCH WITH FIG. 1B

FIG. 1B

MATCH WITH FIG. 1A

360

GGTTGGAGTTGAGTTCCTGCTCTCTGATATTTTAAACGACGTCGTGTAATATATGTC
N L N S R T E E T I K F A A A H Y N T E -

301

C

420

AGATCTTGAAAAAGTATTGATAATGAGTGGAGAAAGACTCAATGCATGCCACGGGAGGTGT
TCTAGAACTTTTCATAACTATTACTCACCTCTTTCTGAGTTACGTACGGTGCCTCCACA
I L K S I D N E W R K T Q C M P R E V C -

361

C

480

GTATAGATGTGGGAAGGAGTTTGGAGTCGCGACAAACACCTTCTTTAAACCTCCATGTG
CATATCTACACCCCTTCCTCAAACCTCAGCGCTGTTTGTGGAAGAAATTTGGAGGTACAC
I D V G K E F G V A T N T F F K P P C V -

421

C

540

TGTCGGCTACAGATGTGGGGTTGCTGCAATAGTAGGGGCTGCAGTGCATGAACACCA
ACAGGCAGATGTCTACACCCCCAACGACGTTATCATTCCCGACGTCACGTACTTGTGGT
S V Y R C G C C N S E G L Q C M N T S -

481

C

600

GCACGAGCTACCTCAGCAAGACGTTATTGAAATTACAGTGCCTCTCTCAAGGCCCA
CGTGCTCGATGGAGTCGTTCTGCAATAAATTAATGTCACGGAGAGAGATTCCGGGT
T S Y L S K T L F E I T V P L S Q G P K -

541

C

660

AACCAGTAACAATCAGTTTTGCCAATCACACTTCCTGCCGATGCATGCTCTAAACTGGATG
TTGGTCATTGTTAGTCAAAACGGTTAGTGTGAAGACGGCTACGTACAGATTTGACCTAC
P V T I S F A N H T S C R C M S K L D V -

601

C

MATCH WITH FIG. 1C

FIG. 1C

MATCH WITH FIG. 1B

661	TTTACAGACAAGTTCATTCCATTATTAGACGTTCCCTGCCAGCAACACTACCACAGTGTCTC	720
	AAATGCTGTTCAAGTAAGGTAATAATCTGCAAGGGACGGTCTGTGTGATGGTGTACACAG	
C	Y R Q V H S I I R R S L P A T L P Q C Q	
721	AGGCAGCGAACAAGACCTGCCCCACCAATTACATGTGGAATAATCACATCTGCAGATGCC	780
	TCCGTCGCTTGTCTGGACGGGGTGTAAATGATACACCTTATTAGTAGACGCTCTACGG	
C	A A N K T C P T N Y M N N H I C R C L	
781	TGGCTCAGGAAGATTTATGTTTTCCTCGGATGCTGGAGATGACTCAACAGATGGATTCC	840
	ACCGAGTCTCTTAAATAACAAAGGAGCCCTACGACCTCTACTGAGTTGTCTACCTAAGG	
C	A Q E D F M F S S D A G D S T D G F H	
841	ATGACATCTGTGGACCAAAAGAGCTGGATGAAGAGACCTGTCTAGTGTCTGCAGAG	900
	TACTGTAGACACCTGGTTTGTCTCCTCGACCTACTTCTCTGGACAGTCAACACAGACGCTCTC	
C	D I C G P N K E L D E E T C C Q C V C R A	
901	CGGGGCTTCGGCCTGCCAGCTGTGGACCCCAAGAACTAGACAGAAACTCATGCCAGT	960
	GCCCCGAAGCCGGACGGTCGACACCTGGGGTGTCTTCTGTATCTGTCTTTGAGTACGGTCA	
C	G L R P A S C G P H K E L D R N S C Q C	
961	GTGCTCTGTAAAAACAACCTCTTCCCCAGCCAATGTGGGGCCAAACCGAGAAATTTGATGAAA	1020
	CACAGACATTTTGTGTTGAGAGGGGTGGTTACACCCCGGTTGGCTCTTAAACTACTTTT	

MATCH WITH FIG. 1D

FILED

V C K N K L F P S Q C G A N R E F F D E N -

ACACATGCCAGTGTGTATGTATAAAGAACCTGTGCCCCAGAAATCAACCCCTAATAATCCTGGAA
1080

1021
TGTGTACGGTCACACATACATTTCTTGGACGGGTCCTTTAGTTGGGATTAGGACCTT
T C C Q C C V C C K R T C C P R N Q P L N P G K -

AATGTGCTGTGAATGTACAGAAAGTCCACAGAAATGCTTGTTAAAAGGAAGAAGTTCC
-----+-----+-----+-----+-----+-----+
1081

1141 ACCACCAACATGCAGCTGTTACAGACGGCCATGTACGAACCGCCAGAGGCTTGTGAGC + 1200
TGGTGGTTGTACGTCGACAATGTCTGCCGGTACATGCTTGGCGTCTTCGGAACACTCG
H Q T C S C Y R R P C T N R Q K A C E P -

1201 CAGGATTTTCATATAGTGAAGAGTGTCGTTGTGTCCTTCATATATGGCAAGACCAC
-----+-----+-----+-----+-----+
1202 GTCCATAAAGTATATCATTCTTCACACAGCAACACAGGAGTATAACCGTTTCTGGTG
G F S Y S E E V C R C V P S Y W Q R P Q -

AAATGAGCTAAGATTGTACTGTTTCCAGGTCATCGATTTTCTATTATGGAACCTGTGT

MATCH WITH FIG. 1E

U

*
S
M

1320

TTTACTCGATTCTTAACATGACAAAAGGTCAAGTAGCTAAAAGATAATACCTTTTGACACA

1261

TGCCACAGTAGAACTGTCCTGTGAACAGAGAGAGACCCCTTGTGGGTCATGCTAAACAAGACA 1380

ACGGTGTTCATCTTGACAGACACTTGTCTCTCTGGGAACACCCAGGTACGATTGTTTCTGT

1321

AAAGTCGTCTTTTCTCGAACCATGTGGATAACCTTACAGAAATGGACCTGGACCTGACCTCGAGTAGAC 1440

CAAAAGGCTCTGTAAAGAGCTGGTTTCTGCCAATGACCAACAGCCAAGATTTTCCTC
-----+-----+-----+-----+-----+-----+-----+-----+ 1500
GTTTTCGGAGAACATTTCTGACCAAAAGACGGTTACTGGTTTGTCCGGTCTTAAAGGAG

TTGTGATTTCTTTAAAAGGATGACTATATATATATTTTCCACTAAAAATATTTGTTCTGC
+-----+-----+-----+-----+-----+
AACACTAAAGAAATTTTCTTACTGATATATTTAAATTAAGGTGATTTTATATACAAAGACG
1560

ATTCATTTTATAGCAACAACAAATTGGTA AA ACTCCTGTGATCAATA TTTTATATATCAT
-----+-----+-----+-----+-----+-----+-----+
TAAGTAAAAATATCGTTGTTTGTTAACCA TTTTGAGTGCACACTAGTTATATAA AATATAGTA

GC AAAATATGTTTAAATAAAAAATGAAAATCTGTATTATTATAAAAAAAA
 1674
 CGTTTTATACAAAAATTTTATTTTACTTTTAAACATAAATATTTT

```

1  CGAGGCCAGGCTTATGCAAGCAAAGATCTGGAGGAGCAGTTACGGTCTGTGTCCAGTGT
   -----+-----+-----+-----+-----+-----+-----+-----+
71  AGATGAACATCATGACTGTACTCTACCCAGAAATATTGGAAAAATGTACAAGTGTACAGCTAAG
   -----+-----+-----+-----+-----+-----+-----+-----+
      M T V L Y P E Y W K M Y K C Q L R
   -----+-----+-----+-----+-----+-----+-----+-----+
121 GAAAGGAGCGTGGCAACATAACAGAGAAACAGGCCAACCTCAACTCAAGGACAGAAAGAGAC
   -----+-----+-----+-----+-----+-----+-----+-----+
      K G G W Q H N R E Q A N L N S R T E E T
   -----+-----+-----+-----+-----+-----+-----+-----+
181 TATAAAATTGCTGCAGCACATTATAATACAGAGATCTTGAAAAGTATTGATAATGAGTGT
   -----+-----+-----+-----+-----+-----+-----+-----+
      I K F A A A H Y N T E I L K S I D N E W
   -----+-----+-----+-----+-----+-----+-----+-----+
241 GAGAAAGACTCAATGCATGCCACGGGAGGTGTGTATAGATGTGGGAAGGAGTTTGGAGT
   -----+-----+-----+-----+-----+-----+-----+-----+
      R K T Q C M P R E V C I D V G K E F G V
   -----+-----+-----+-----+-----+-----+-----+-----+
301 CGGACAAACACCTTCTTTAAACCTCCATGTGTGTCCGTCTACAGATGTGGGGGTTGCTG
   -----+-----+-----+-----+-----+-----+-----+-----+
      A T N T F F K P P C V S V Y R C G G C C
   -----+-----+-----+-----+-----+-----+-----+-----+

```

FIG. 2A

```

361 CAATAGTGGGGCTGCAGTGCATGAACACCAGCAGCTACCTCAGCAAGACGTTATT
-----+-----+-----+-----+-----+
   N S E G L Q C M N T S T S Y L S K T L F

421 TGAAATTACAGTGCCTCTCTCTCAAGGCCCCAAACCAGTAACAATCAGTTTGGCCAATCA
-----+-----+-----+-----+-----+
   E I T V P L S Q G P K P V T I S F A N H

481 CACTTCCTGCCGATGCATGCTCTAACTGGATGTTTACAGACAAAGTTCCATTCCATTATTAG
-----+-----+-----+-----+-----+
   T S C R C M S K L D V Y R Q V H S I I R

541 ACGTTCCTGCCAGCAACACTACCACAGTGTCTAGGCAGCGAACAAGACCTGCCCCACCAA
-----+-----+-----+-----+-----+
   R S L P A T L P Q C Q A A N K T C P T N

601 TTACATGTGGAATAATCACATCTGCAGATGCCTGGCTCAGGAAGATTTATGTTTTCCTC
-----+-----+-----+-----+-----+
   Y M W N N H I C R C L A Q E D F M F S S

661 GGATGCTGGAGATGACTCAACAGATGGATTCCATGACATCTGTGGACCAACAAGGAGCT
-----+-----+-----+-----+-----+
   D A G D D S T D G G F H D I C G P N K E L

```

FIG.2B

```

721 GGATGAAGAGACCTGTAGTGTCTGCAGAGGGGGCTTCGGCCTGCCAGCTGTGGACC
    D E E T C Q C V C R A G L R P A S C G P

781 CCACAAAGAACTAGACAGAACTCATGCCAGTGTGTCTGTATAAAACAACTCTTCCCCAG
    H K E L D R N S C Q C V C K N K L F P S

841 CCAATGTGGGCCCAACCGAGAAATTGATGAAACACATGCCAGTGTGTATGTAAGAAGAAC
    Q C G A N R E F D E N T C Q C V C K R T

901 CTGCCCCAGAAATCAACCCCTAAATCCTGGAAATGTGCCCTGTGAATGTACAGAAAGTCC
    C P R N Q P L N P G K C A C E C T E S P

961 ACAGAAATGCTTGTATAAAGGAAAGAAAGTTCCACCACCAACATGCAGCTGTACAGACG
    Q K C L L K G K K F H H Q T C S C Y R R

1021 GCCATGTACGAACCGCCAGAGGCTTGTGAGCCAGGATTTTCATATAGTGAAGAAGTGTG
    P C T N R Q K A C E P G F S Y S E E V C

```

FIG.2C

1081	TCGTTGTGTCCTTCATATTGGCAAAGACACCACAAATGAGCTAAGATTGTACTGTTTCCCA
	R C V P S Y W Q R P Q M S
1141	GTTCAATCGATTTTCTATTATGGAAAACACTGTGTGCCACAGTAGAACTGTCTGTGTAACAGA
1201	GAGACCCCTGTGGGTCCATGCTAACAAAGACAAAAGTCTGTCTTTCCCTGAACCATGTGGA
1261	TAACTTTACAGAAATGGACTGGAGCTCATCTGCAAAAGGCCCTCTTGTAAAGACTGGTTTT
1321	CTGCCAATGACCAAACAGCCAAGATTTTCCCTCTGTGTGATTTCTTTAAAGAATGACTATA
1381	TAATTTATTCCACTAAAAATATTGTTTCTGCATTCATTTTATAGCAACAACAATTGGT
1441	AAAACCTCACTGTGATCAATATTTTATATCATGCAAAATATGTTTAAATAAAATGAAAAA
1501	TTGTATTATAAAAAAAAAAAAAAA

FIG. 2D

50

1

pdgfa .MRTLACLLL LCGCYLAHVL AEEAEIPREV IERLARSQIH SIRDLORLLE
 pdgfb MNRCAW.A.LFL SLCCYLRIVS AEGDPIPEEL YEMLSHSIR SFDDLQRLLEH
 vegfMNFLL SWVHWSLALL LY.....
 vegf2MTV LYPEYWKMYK CQ.....

100

51

pdgfa IDSVGSEDSL DTSIRAHGVH ATKHVPEKRP LPIRRKRSI.EEAVP
 pdgfb GDP.GEEDGA ELDLNMTRSH SGGELES... .LARGRRSLG SLTIAEPAMI
 vegf APMAE..... GGGQ NHHEVVKEFD .VYQR.....
 vegf2 REQANLNSRT EETIKFAAAH YNTEILKSID NEWRK.....

150

101

pdgfa AVCKTRTVIY EIPRSQVDPT SANFLIWPPC VEVKRCCTGCC NTSSVRCQPS
 pdgfb AECKTRTEVF EISRRLLDRT NANFLVWPPC VEVQRCSGCC NNRNVQCRPT
 vegf SYCHPIETLV DIFQEYFDEI ..EYIFKPS VPLMRCGGCC NDEGLECVPT
 vegf2 TQCMPREVCI DVGKEFGVAT ..NTFFKPPC VSVYRCGGCC NSEGLQCMNT

200

151

pdgfa RVHRSVKVA KVEYVRKKPK LKEVQVRLEE HLECAC..... AT.....
 pdgfb QVQLRPVQVR KIEIVRKKPI FKKATVTLED HLACKC.... ETVAARPVT
 vegf EESNITMQIM RIK.PH..QG QHIGEMSFQ HNKCECRPKK DRARQEKKS
 vegf2 STSYLSKTLF EIT.VPLSQG PKPVTISFAN HTSCRCMSKL DVYRQVHSII

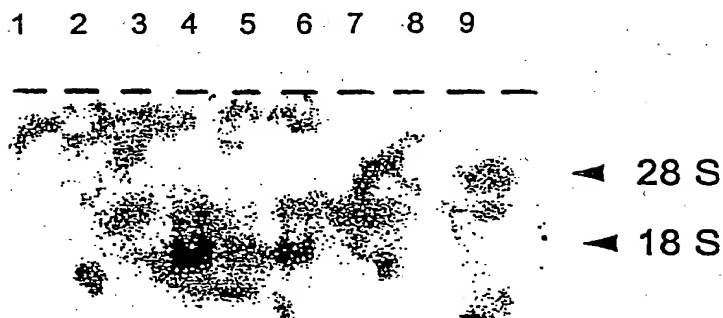
FIG. 3A

201		250
PdgfaTSLNPD YREEDTDVR.
Pdgfb	RSPGGSQEQR AKTPQTRVTI RTVRVRRPPK GKHRKFKKTH DKTALKETLG	
Vegf	RGK.....GKGQKRRK KSRKSWSVY VGARCCIMPW SLPGPHP...	
Vegf2	RRSLPATLPQ CQAANKTCPT NYMWNHICR CLAQEDFMFS SDAGDDSTDG	
251		300
Pdgfa
Pdgfb	A.....
VegfCGP.....CSE RRKHLFVQDE QTCKCSCKNT	
Vegf2	FHDICGPNKE LDEETCQVC RAGLRPASC GPHKEL...DR NSCQCVCKNK	
301		350
Pdgfa
Pdgfb
Vegf	..DSRCKARQ LEINERTC RC DKPRR
Vegf2	LFPSQCGANR .EFDENTCQC VCKRTCPRNQ PLNPGKCACE CTESPQKCLL	
351		398
Pdgfa
Pdgfb
Vegf
Vegf2	KGKKFHHQTC SCYRRPCTNR QKACEPGFSY SEEVRCVPS YWQRPQMS	

PERCENTAGE (%) OF AMINO ACID IDENTITIES BETWEEN EACH PAIR OF GENES IS SHOWN IN THE FOLLOWING TABLE				
	PDGF α	PDGF β	VEGF	VEGF2
PDGF α				
PDGF β	48.0			
VEGF	20.7	22.7		
VEGF2	23.5	22.4	30.0	

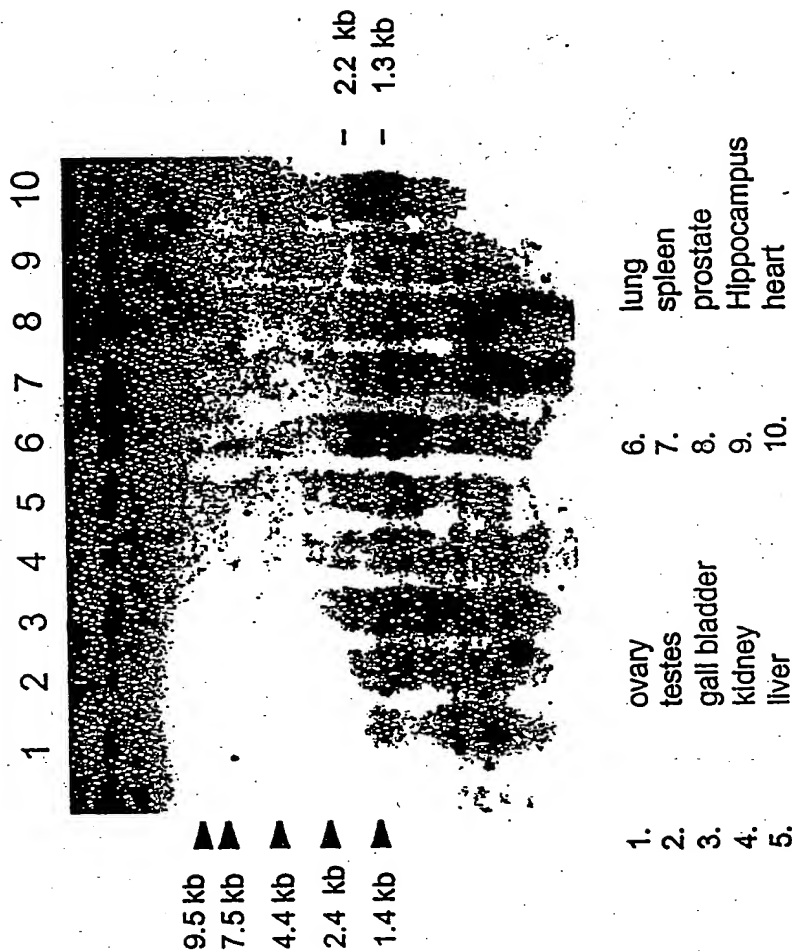
FIG. 4

Expression of VEGF2 mRNA in
Human Breast Tumor Cells



1. normal breast tissue
2. breast tumor tissue
- 3-9. breast tumor cell lines.

FIG. 5

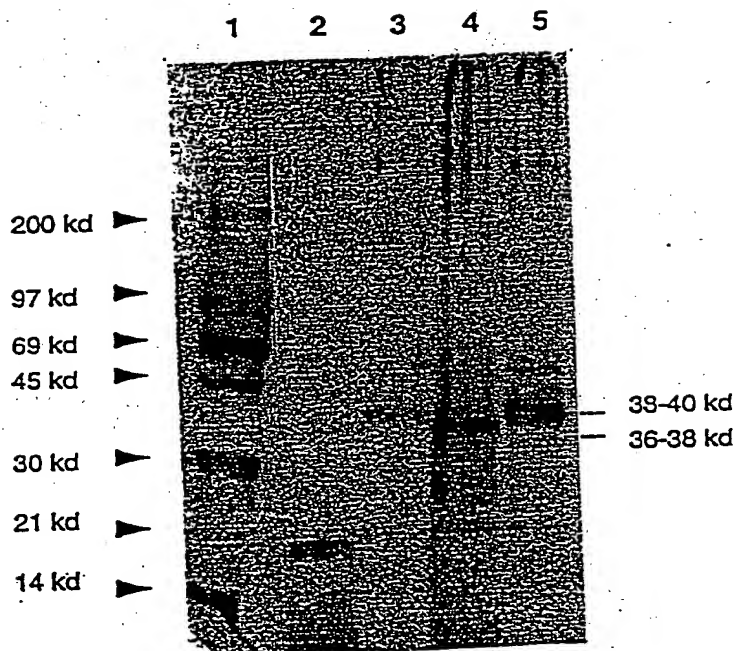


Expression of VEGF2 mRNA in human adult tissues.

FIG. 6

BEST AVAILABLE COPY

FIG. 7



- Lane 1: 14-C and rainbow M.W. marker
 Lane 2: FGF control
 Lane 3: VEGF2 (M13-reverse & forward primers)
 Lane 4: VEGF2 (M13-reverse & VEGF-F4 primers)
 Lane 5: VEGF2 (M13-reverse & VEGF-F5 primers)

non-reducing gel



FIG. 8A

reducing gel

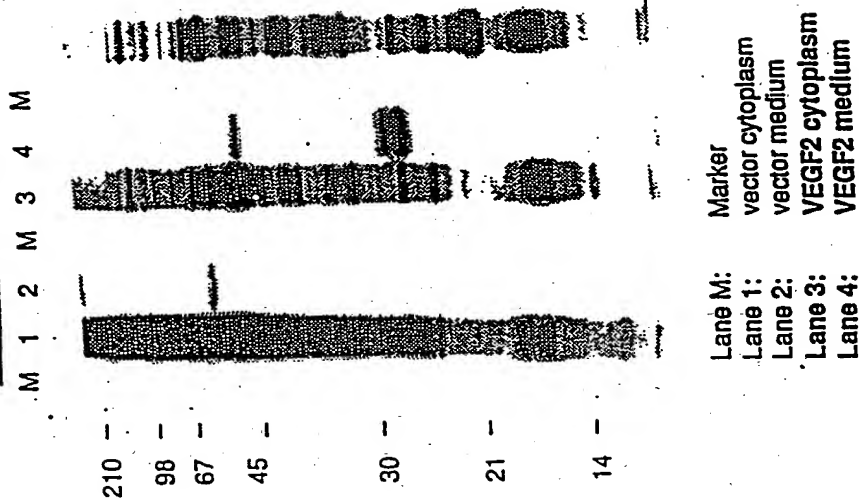
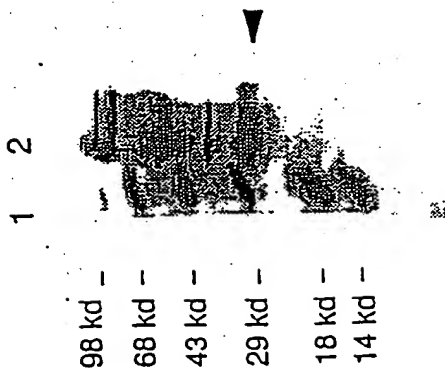


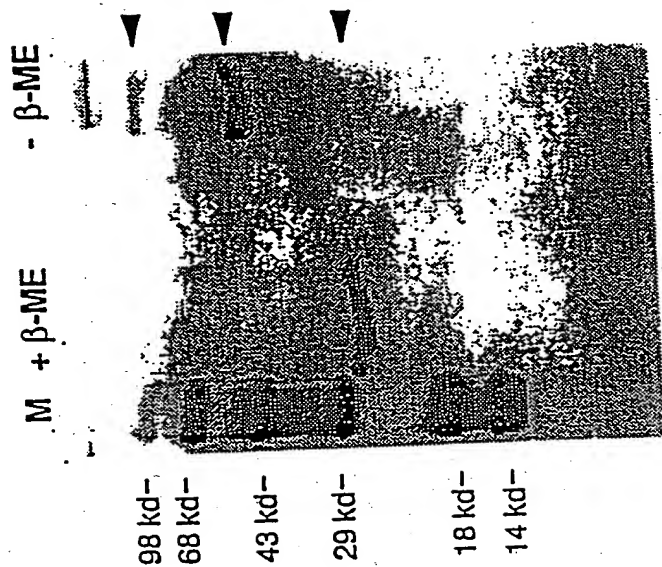
FIG. 8B

FIG. 9



Lane 1: Molecular weight marker
 Lane 2: Precipitates containing VEGF2.

FIG. 10



BEST AVAILABLE COPY

FIG. 11

TD4280-92/SE660

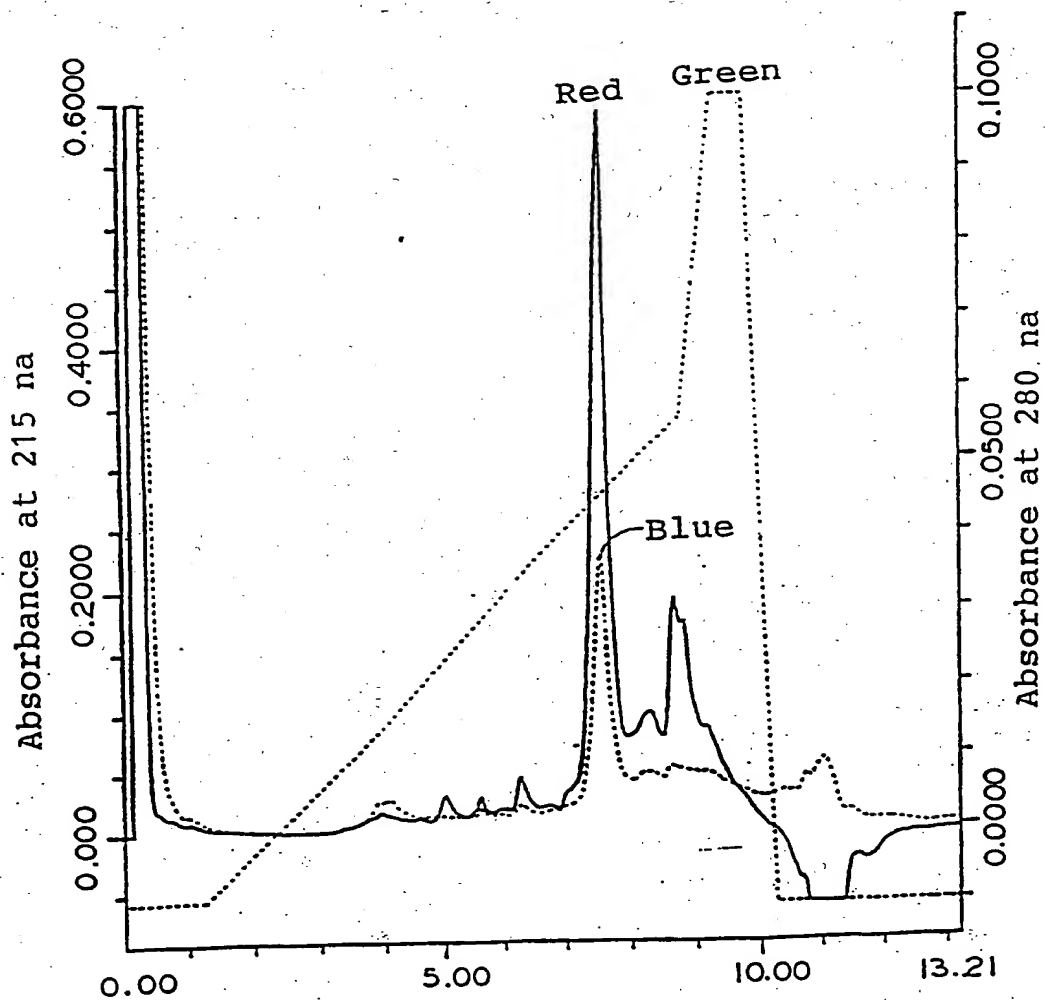


FIG. 12

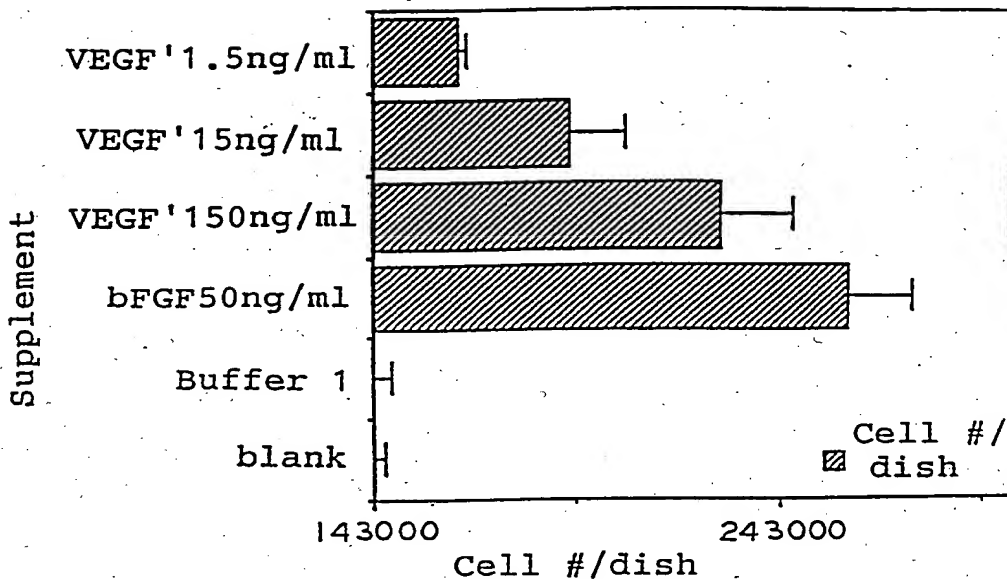
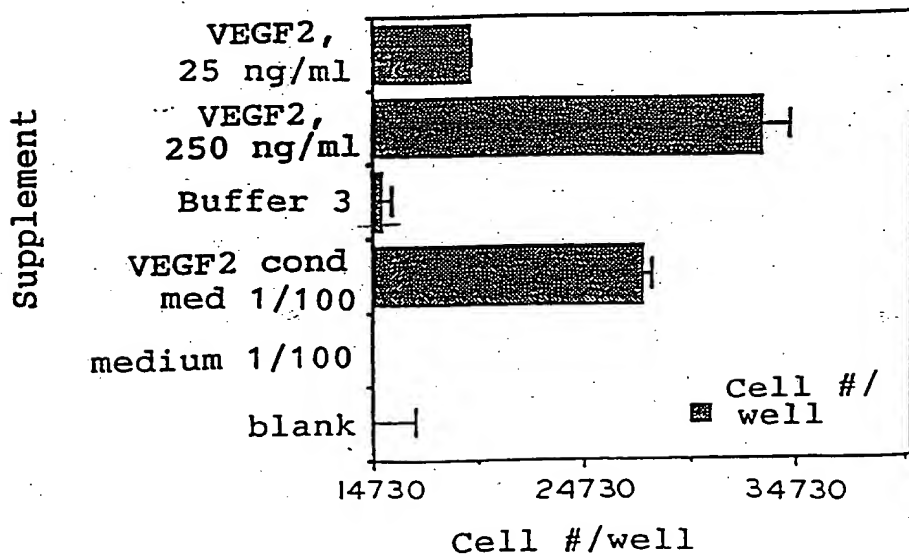


FIG. 13



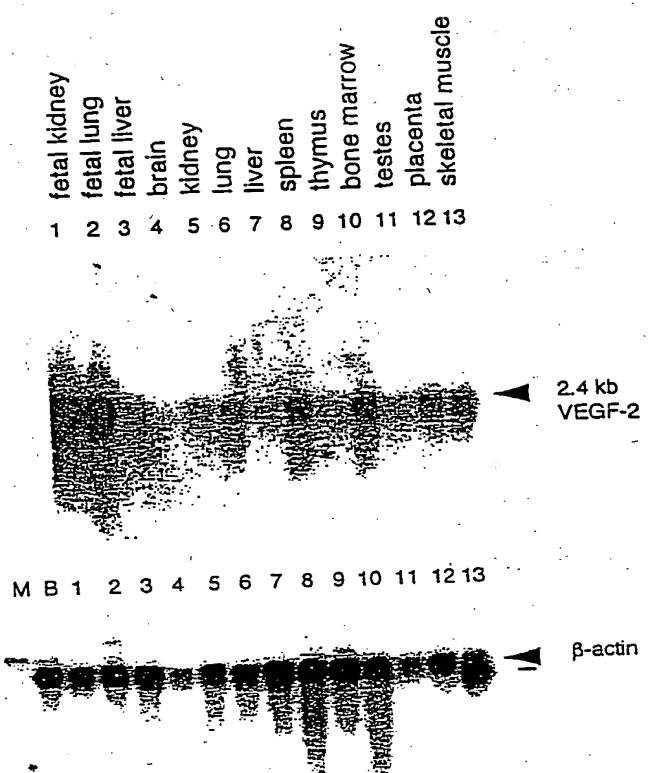


Figure 14

09935726.082401

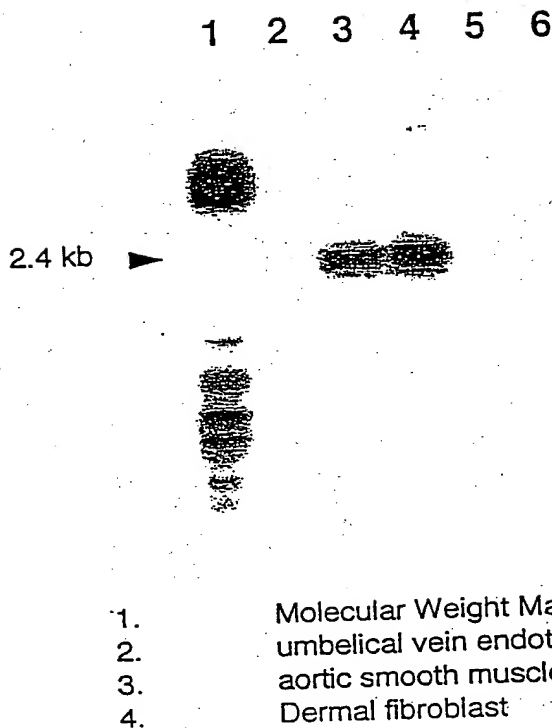


Figure 15

09935726-082401

104230-92/5E660

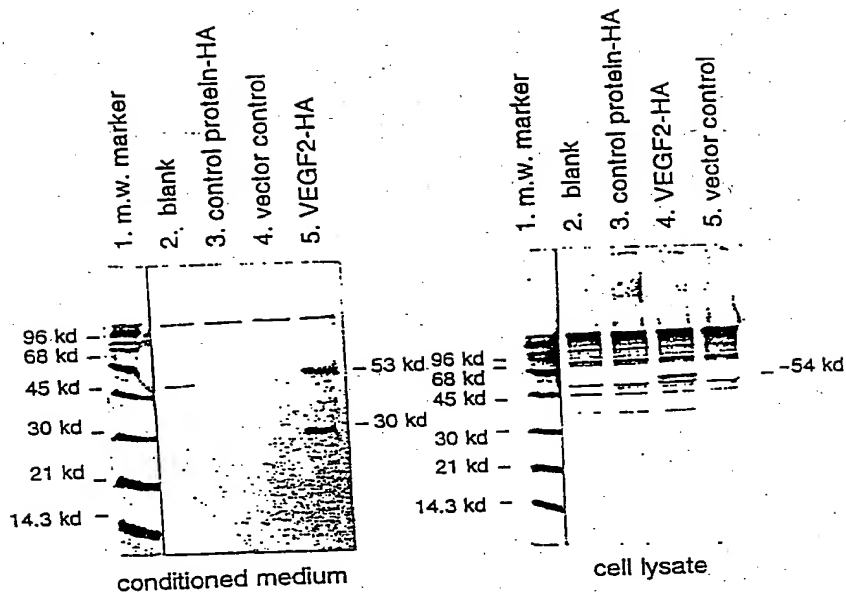


Figure 16

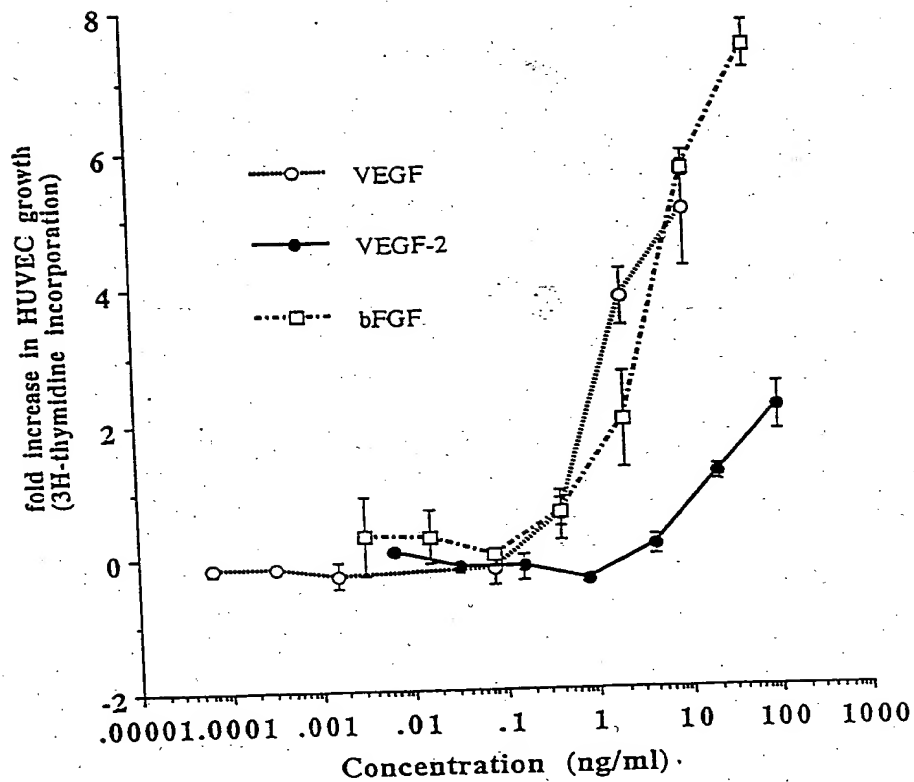


Figure 17

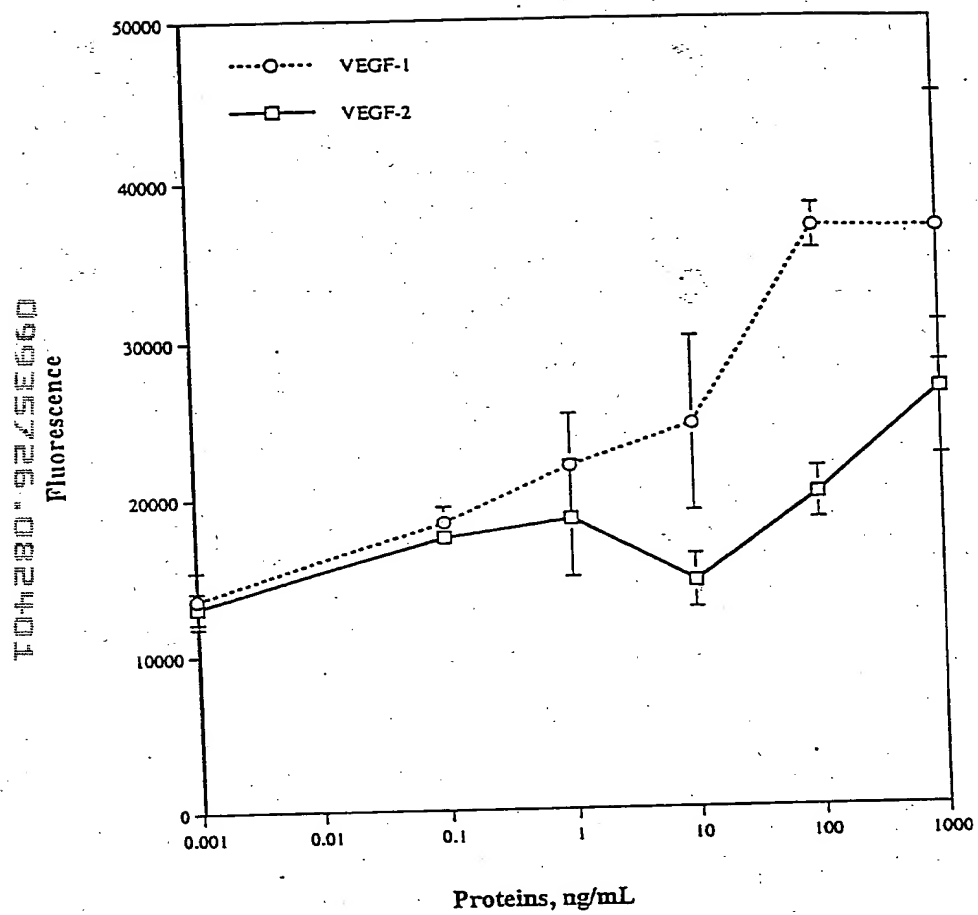


Figure 18

104280-92/5E660

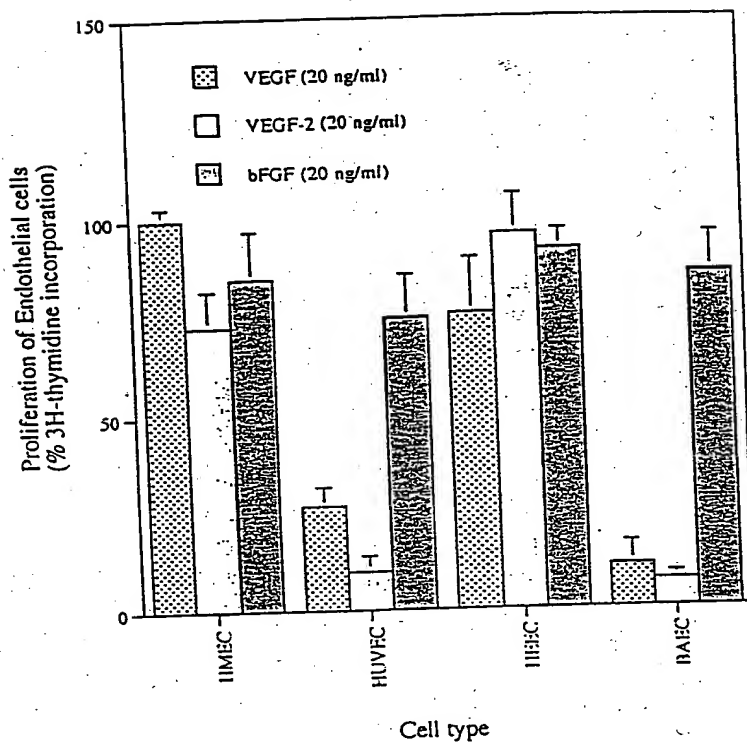
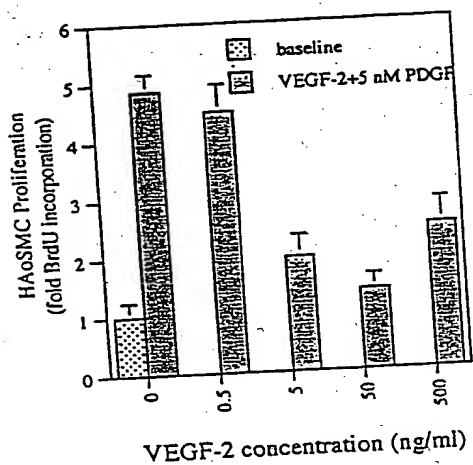


Figure 19

1011280" 9245660

A.



B.

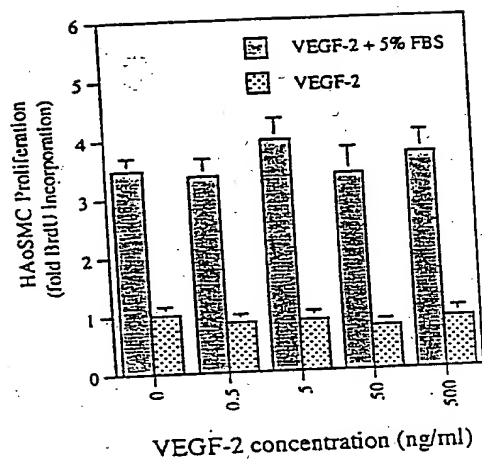
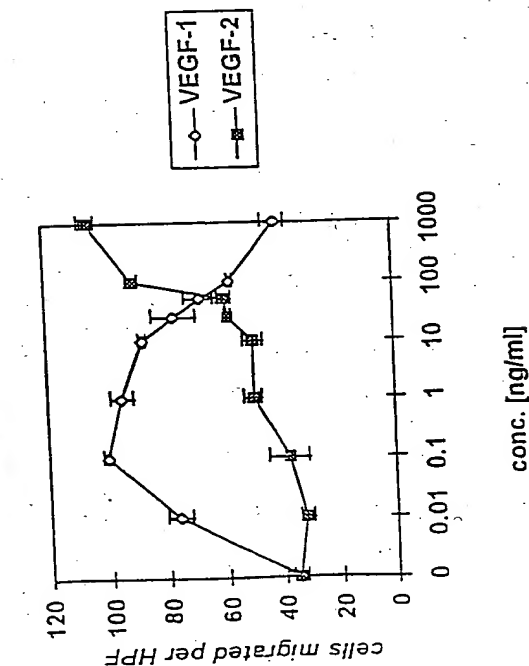


Figure 20

HUVEC Migration



BMEC Migration

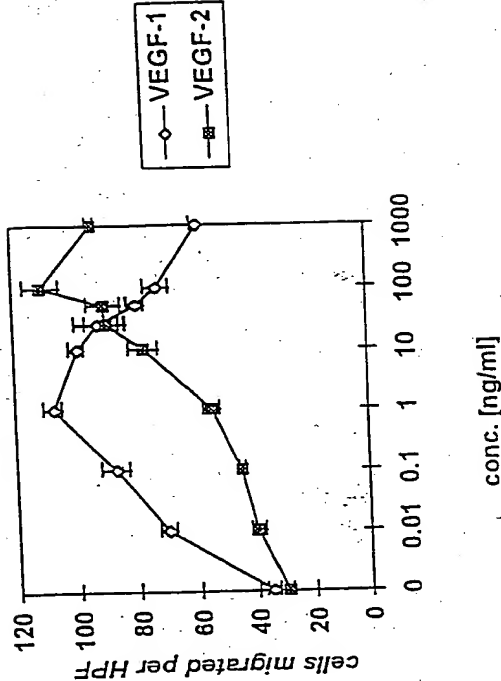


Figure 21

HUVEC - NO-Release

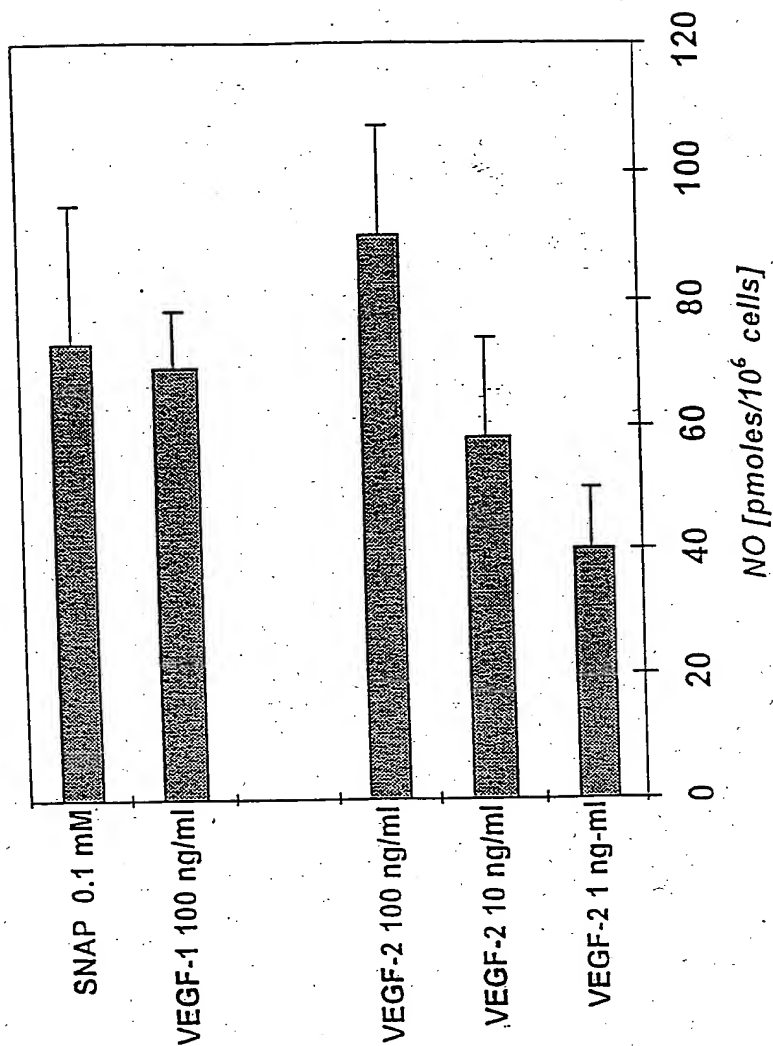


Figure 22

0935726.08401
T04280.9275E60

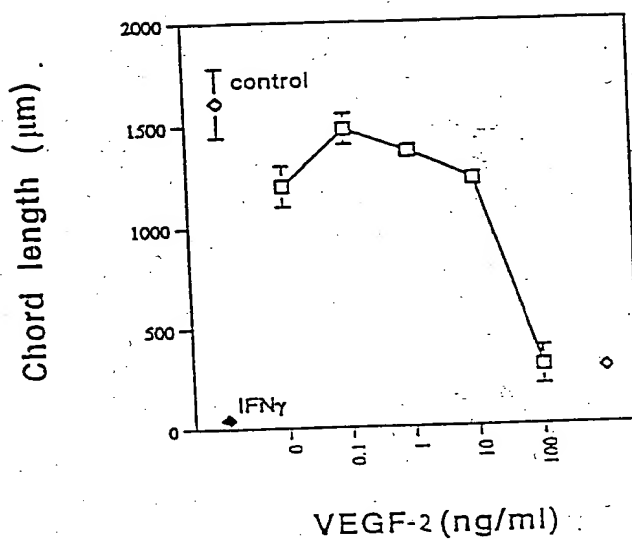


Figure 23

104280-92/5E660

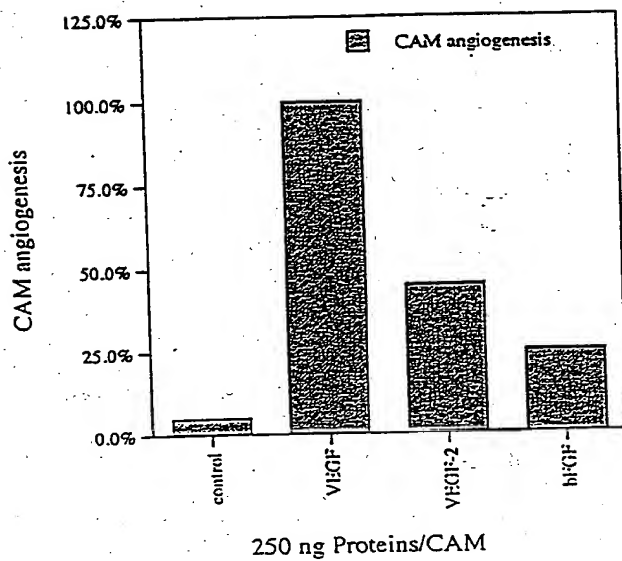


Figure 24

09935726-082401
T04280-9275E660

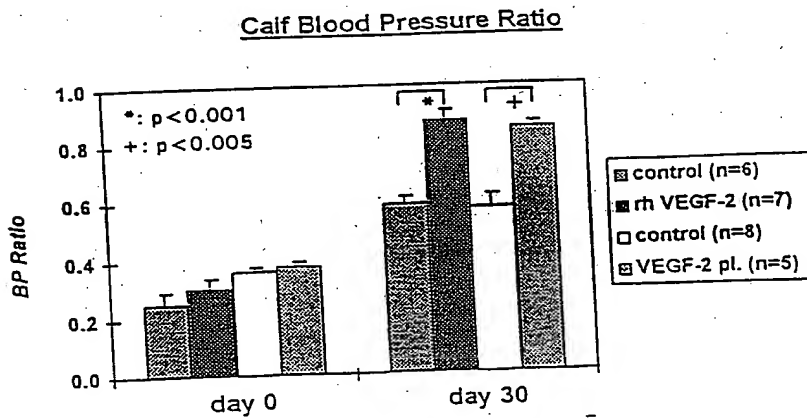
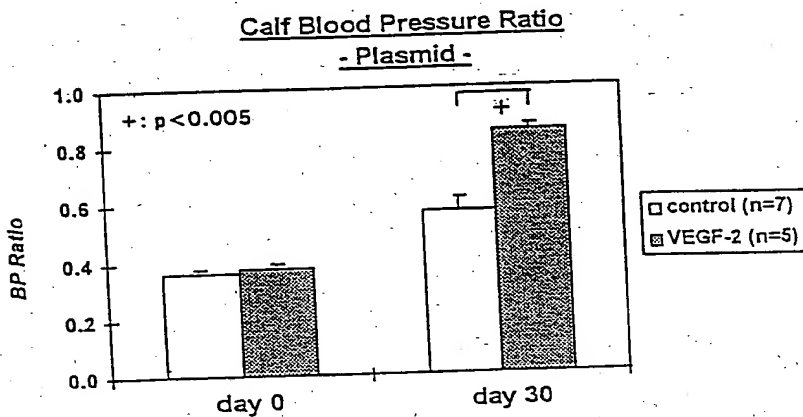
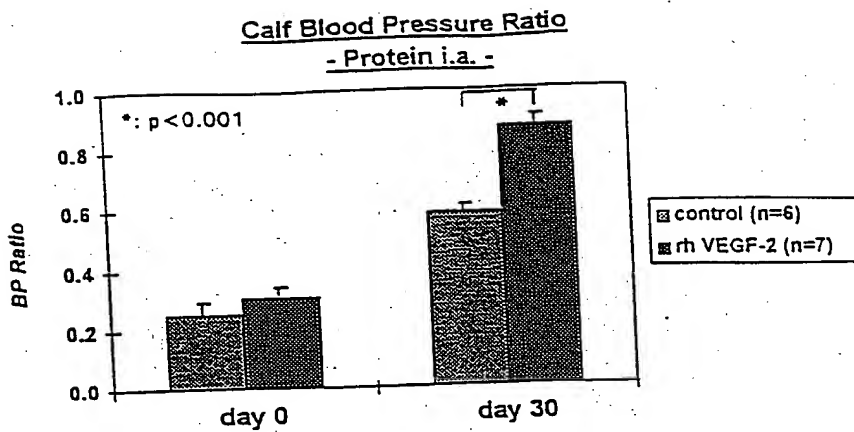


Figure 25A

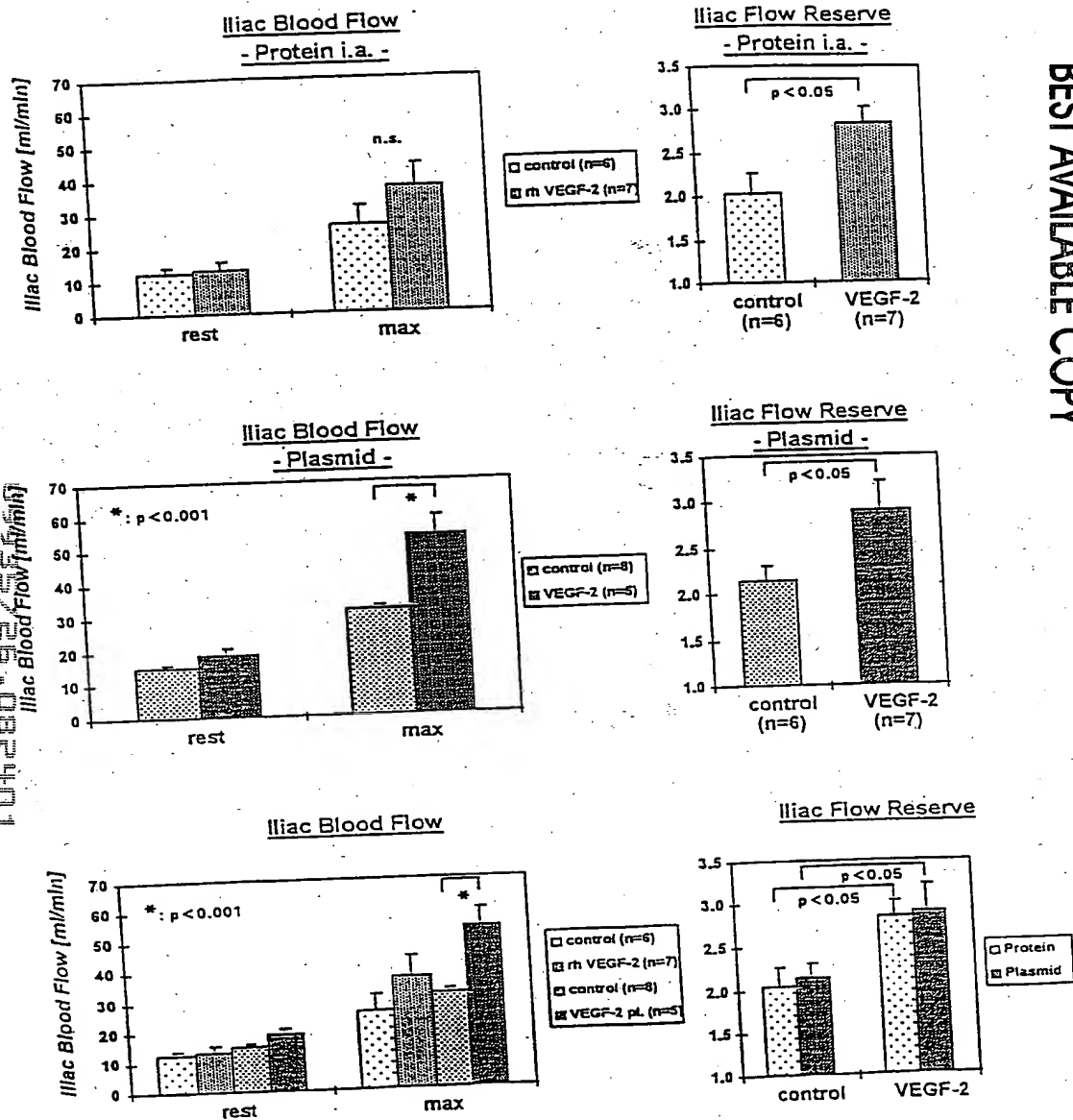
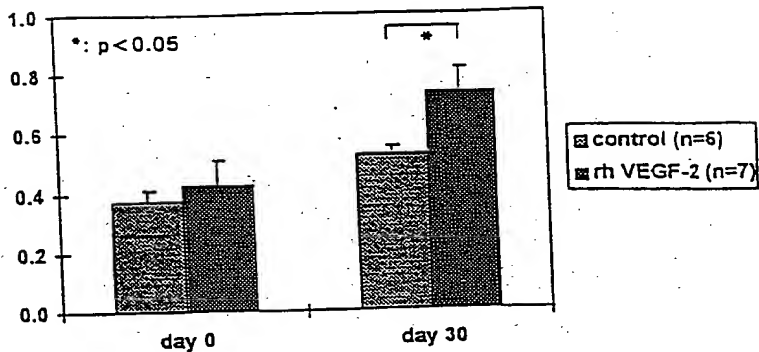


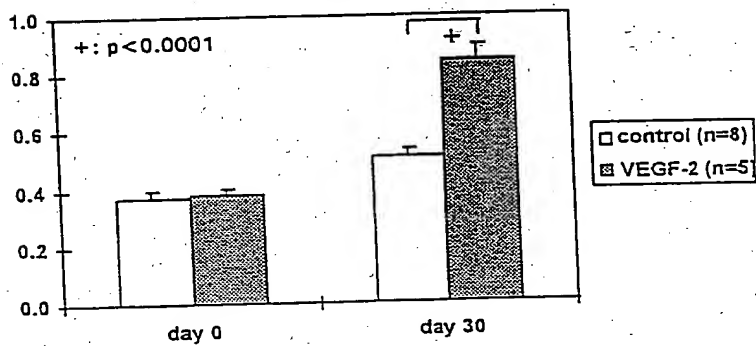
Figure 25B

09935726.082401

Angiographic Score
- Protein i.a. -



Angiographic Score
- Plasmid -



Angiographic Score

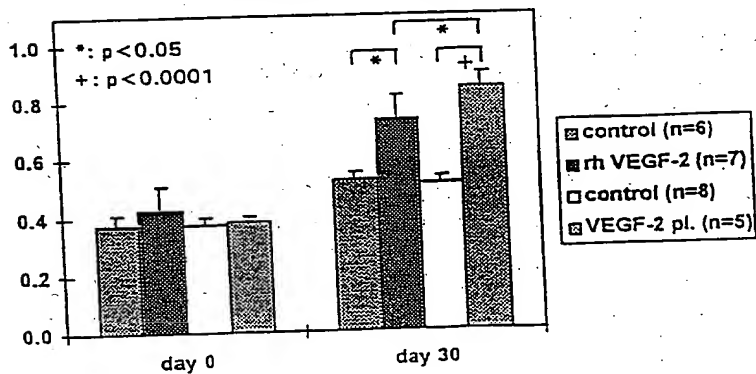


Figure 25C

TD#280" 92/5E660

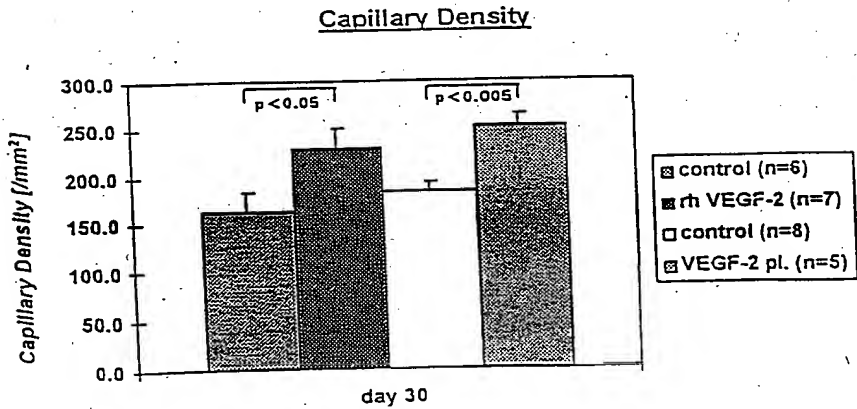
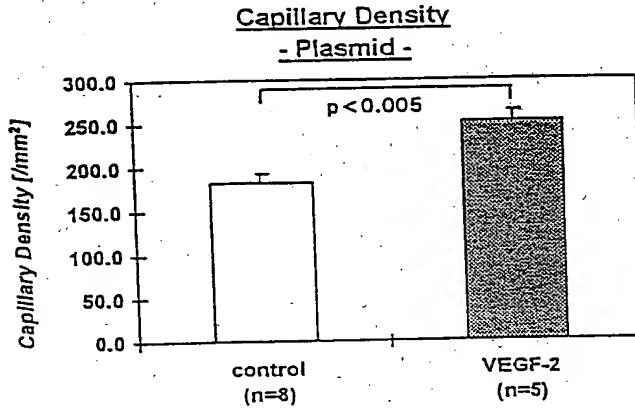
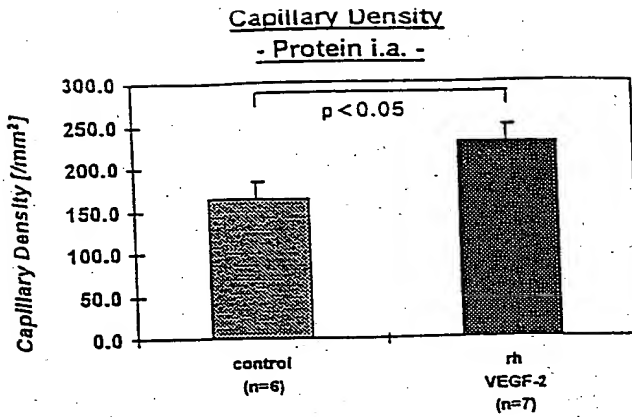


Figure 25D

09955726.082401
T04280" 92/5E660

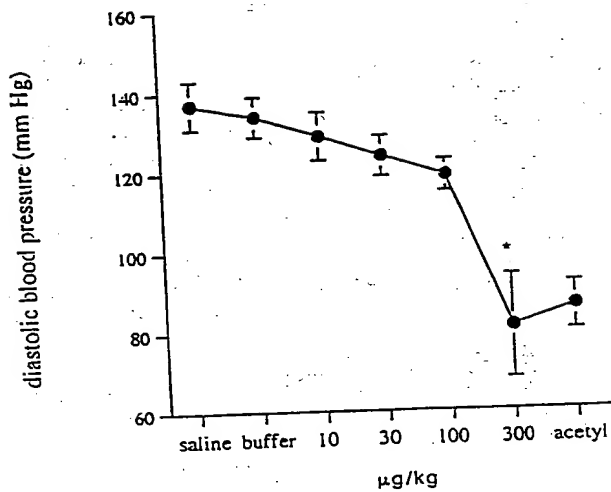


Figure 26A

093576.08401
T04280.9245660

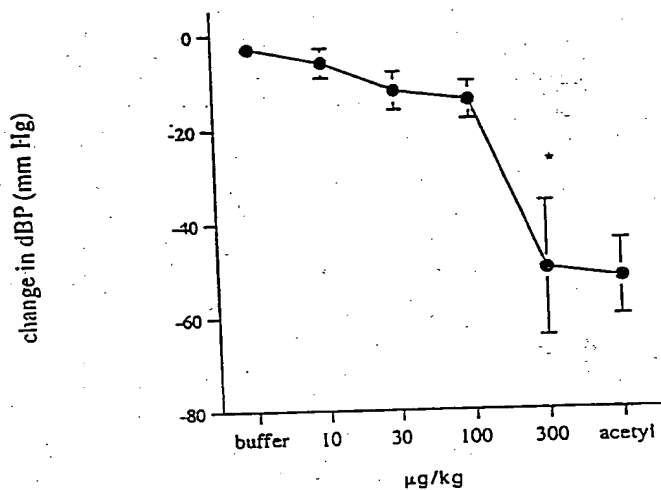


Figure 26B

104280" 9245E660

MAP (mm Hg)

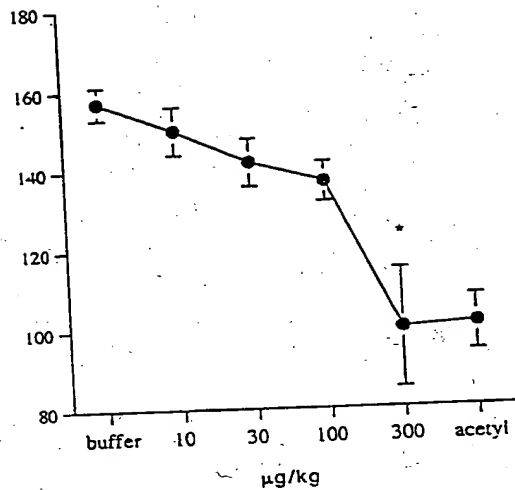


Figure 26C

T04280-9275E660

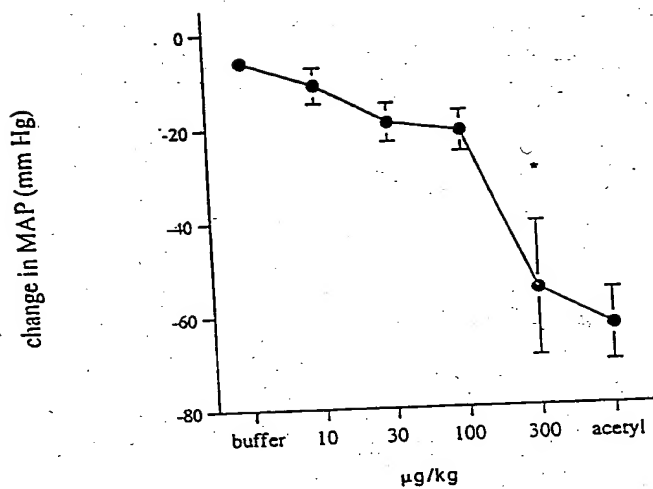
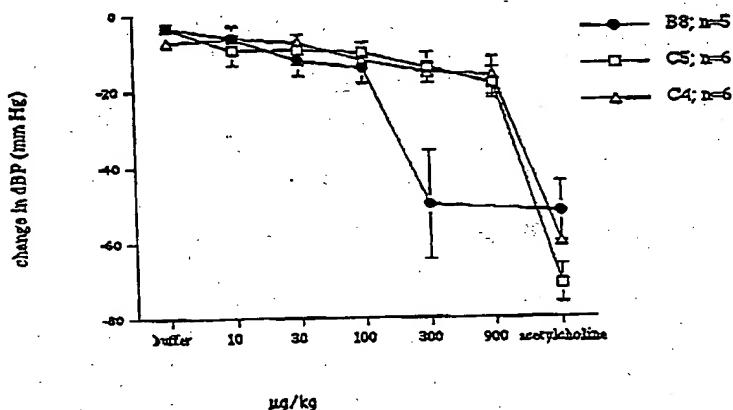


Figure 26D

Change in diastolic blood pressure of SHR rats given increasing doses of VEGF-2

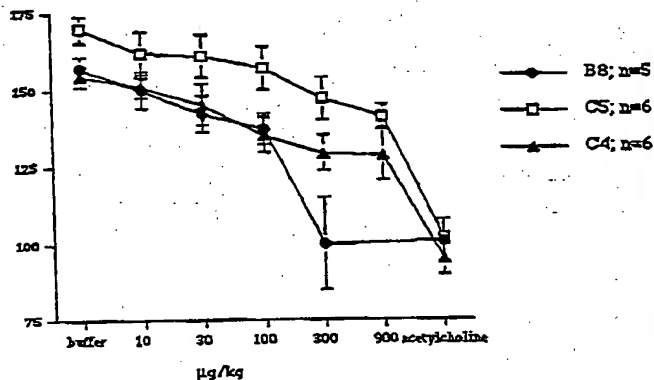


Increasing doses of VEGF-2 (HG00403-B8, HG00404-C5, and HG00404-C4) were administered to 13-14 week old SHR (spontaneously hypertensive rats) and the data are expressed as the mean \pm SEM. Statistical analysis was performed with a paired t-test and significance was defined as $p < 0.05$ vs. the response to buffer alone. The response to B8 and C4 were significant at the 300 μ g/kg dose. The response to C5 was significant at the 100, 300, and 900 μ g/kg doses.

The effect of increasing doses of VEGF-2 on the mean arterial pressure (MAP) of SHR rats

Figure 26E

The effect of increasing doses of VEGF-2 on the mean arterial pressure (MAP) of SHR rats

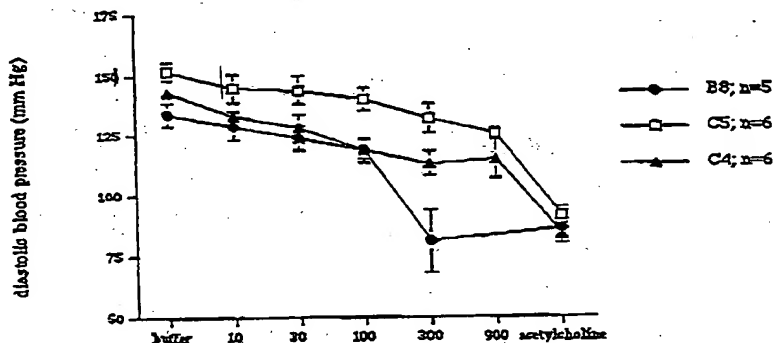


Increasing doses of VEGF-2 (HG00403-B8, HG00404-C5, and HG00404-C4) were administered to 13-14 week old SHR (spontaneously hypertensive rats) and the data are expressed as the mean \pm SEM. Statistical analysis was performed with a paired t-test and significance defined as $p < 0.05$ vs. the response to buffer alone. The response to B8 was significant at a 300 µg/kg dose. Administration of C5 yielded significant responses at doses greater than or equal to 100 µg/kg. The response to C4 was significant when 10, 100, 300, and 900 µg/kg were given.

The effect of VEGF-2 on the diastolic blood pressure of SHR rats

Figure 26F

The effect of VEGF-2 on the diastolic blood pressure of SHR rats



Increasing doses of VEGF-2 (HG00403-B8, HG00404-CS, HG00404-C4) were administered to 13-14 week old SHR (spontaneously hypertensive rats) and the data are expressed as the mean \pm SEM. Statistical analysis was performed with a paired t-test and statistical significance was defined as $p < 0.05$ vs. the response to buffer alone. The response to B8 was significant only at the 300 μ g/kg dose and when given acetylcholine. The responses to C4 and CS, while much less dramatic, were statistically significant at all dose levels.

Figure 26G

104280-9275660

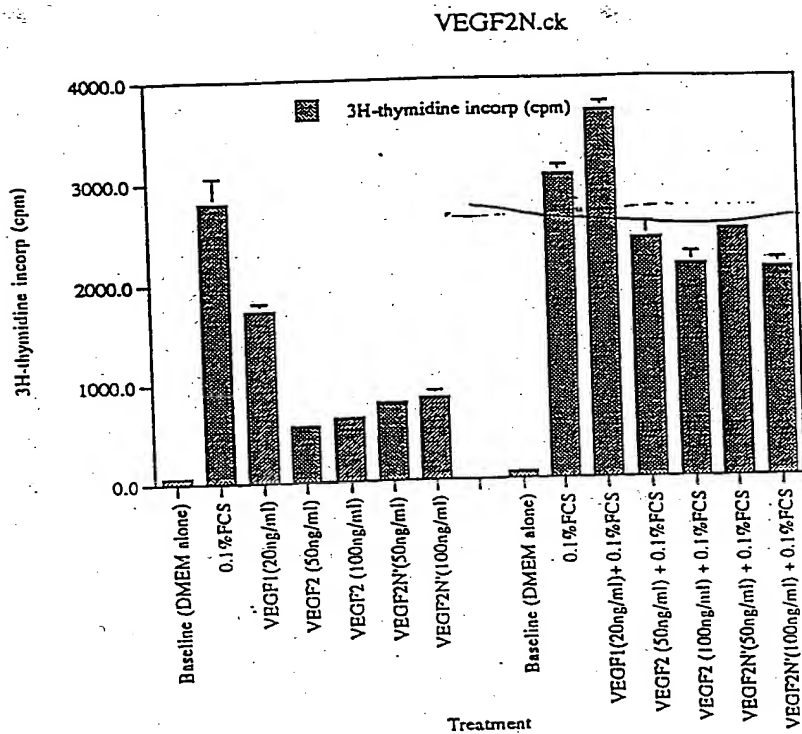


Figure 27

09935726.082401
104280"9225E660

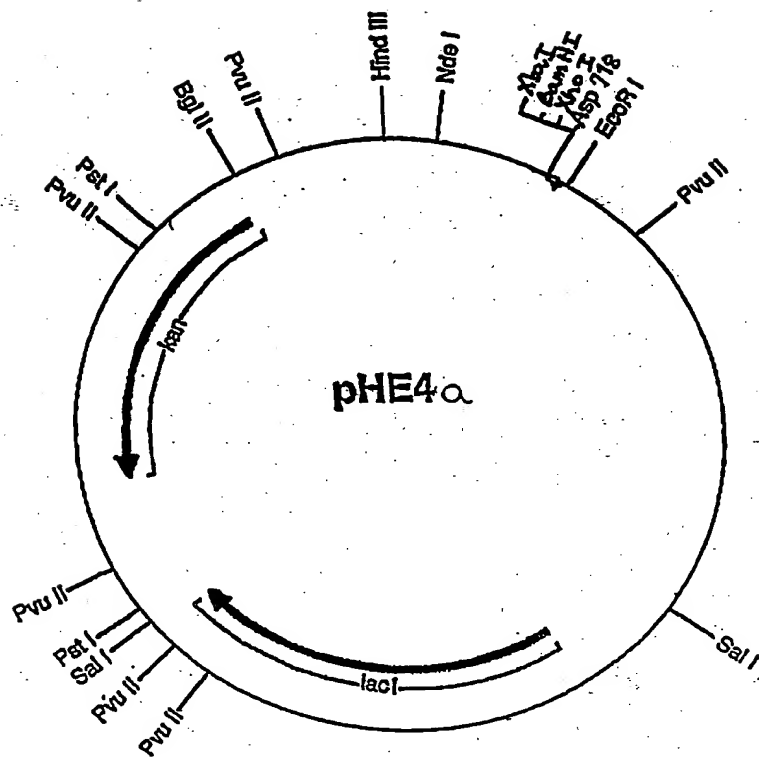


Figure 28

